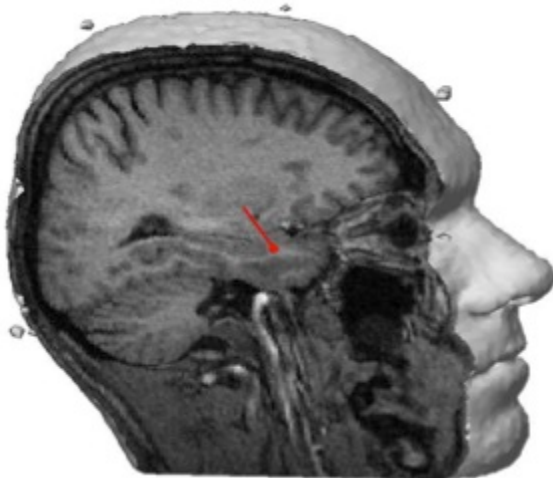
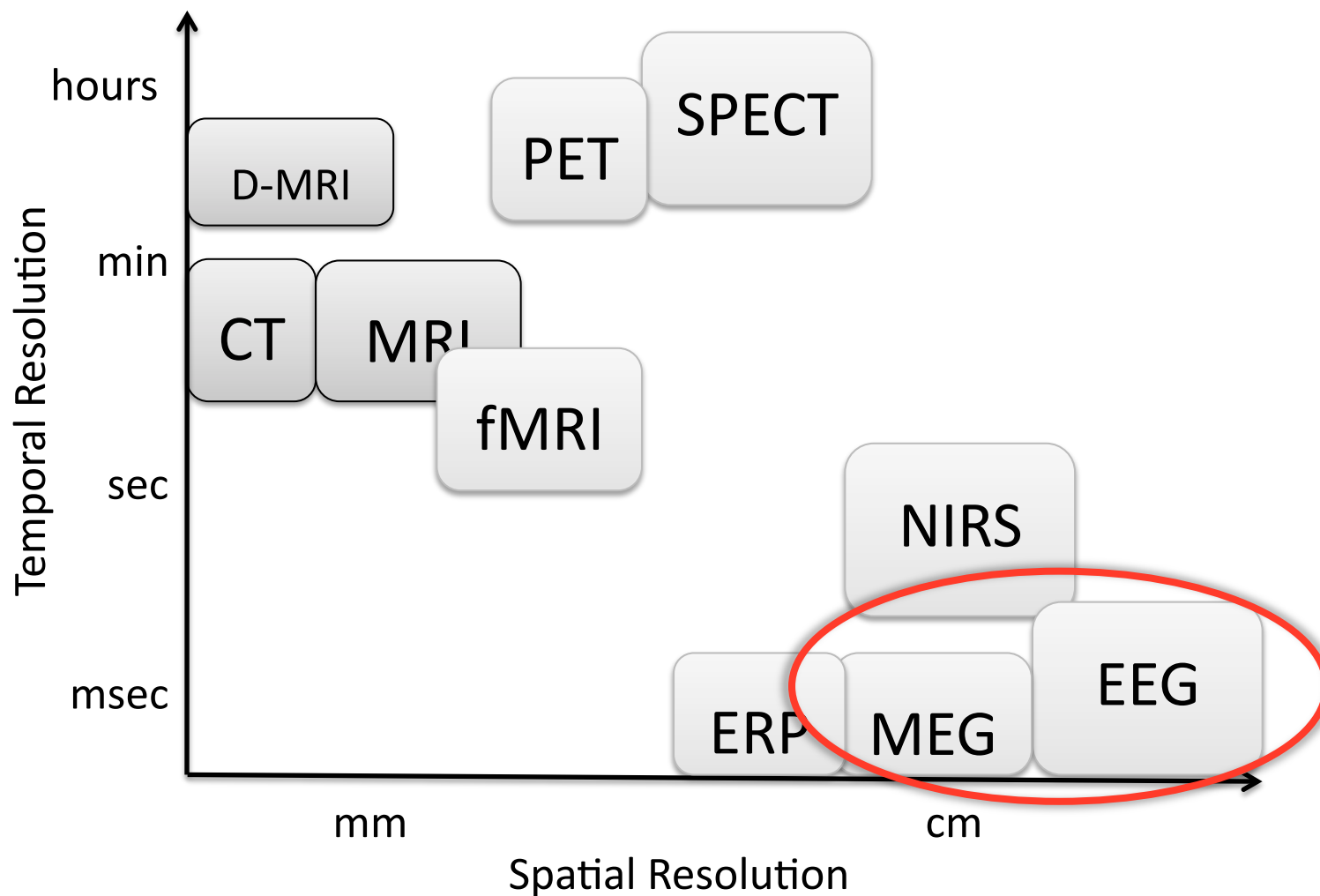


# Brain Imaging through EEG and MEG

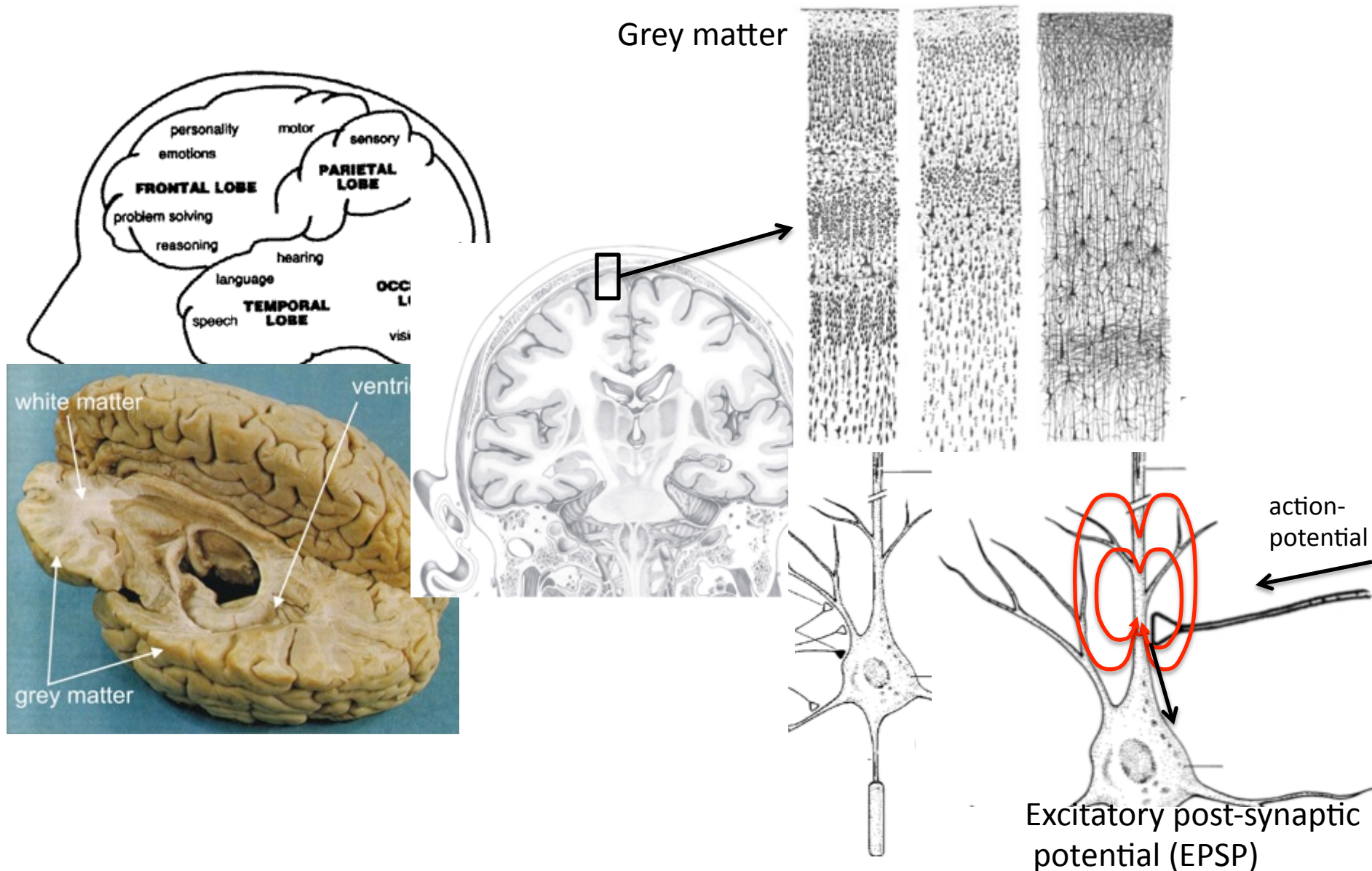
Dr. ir. Hans Hallez



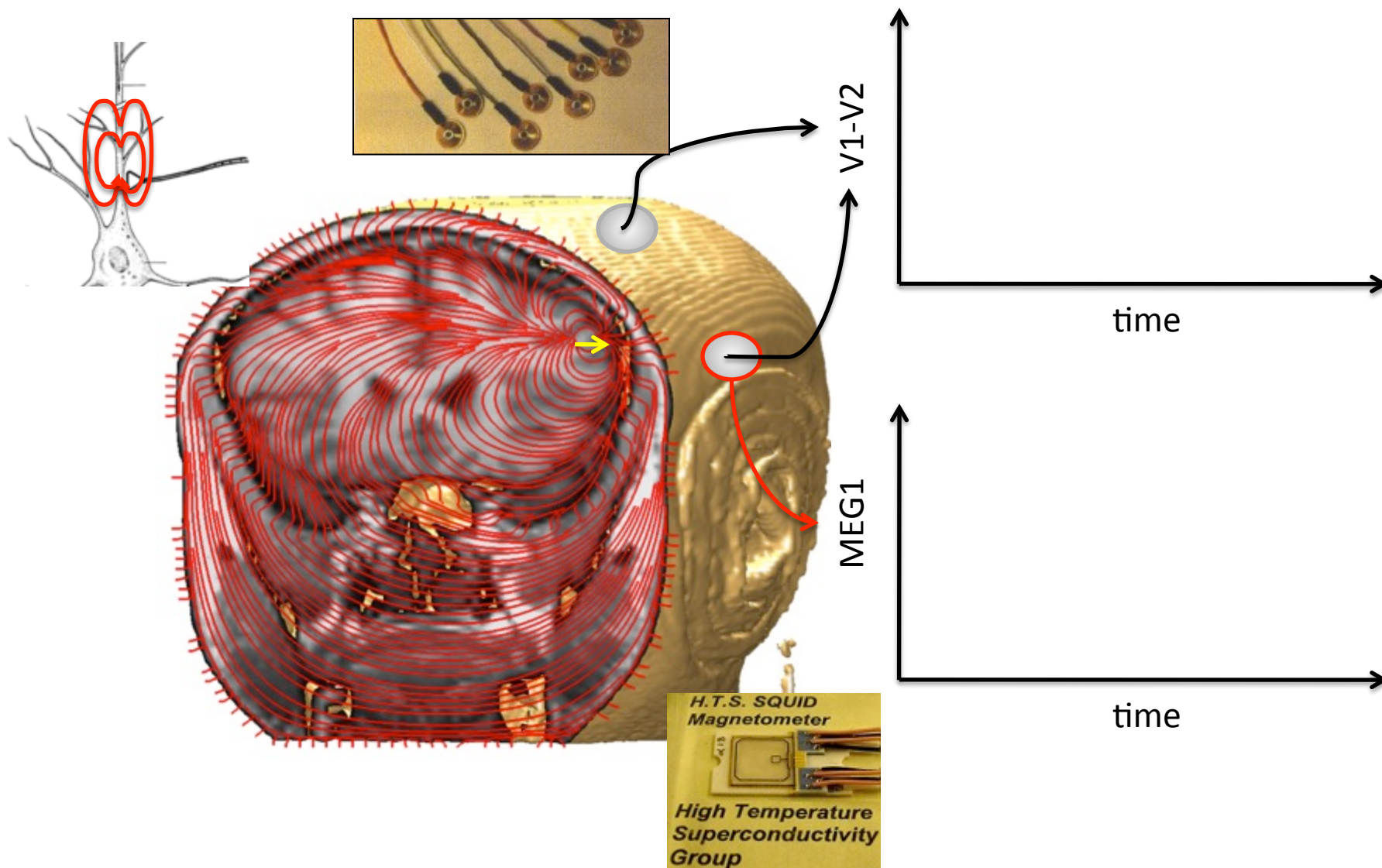
# Imaging the brain



# Neurons as electrical generators

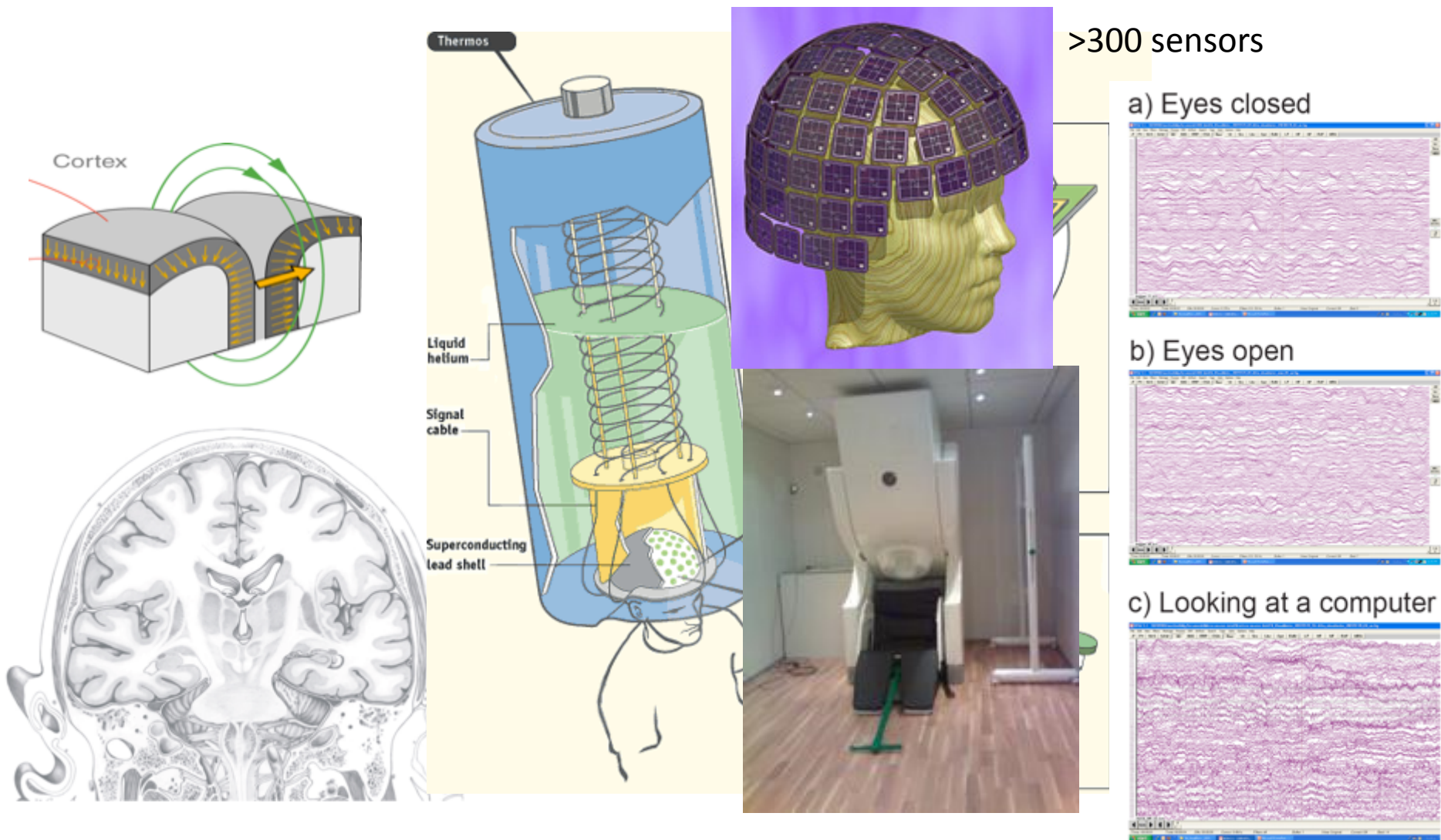


# Volume conduction

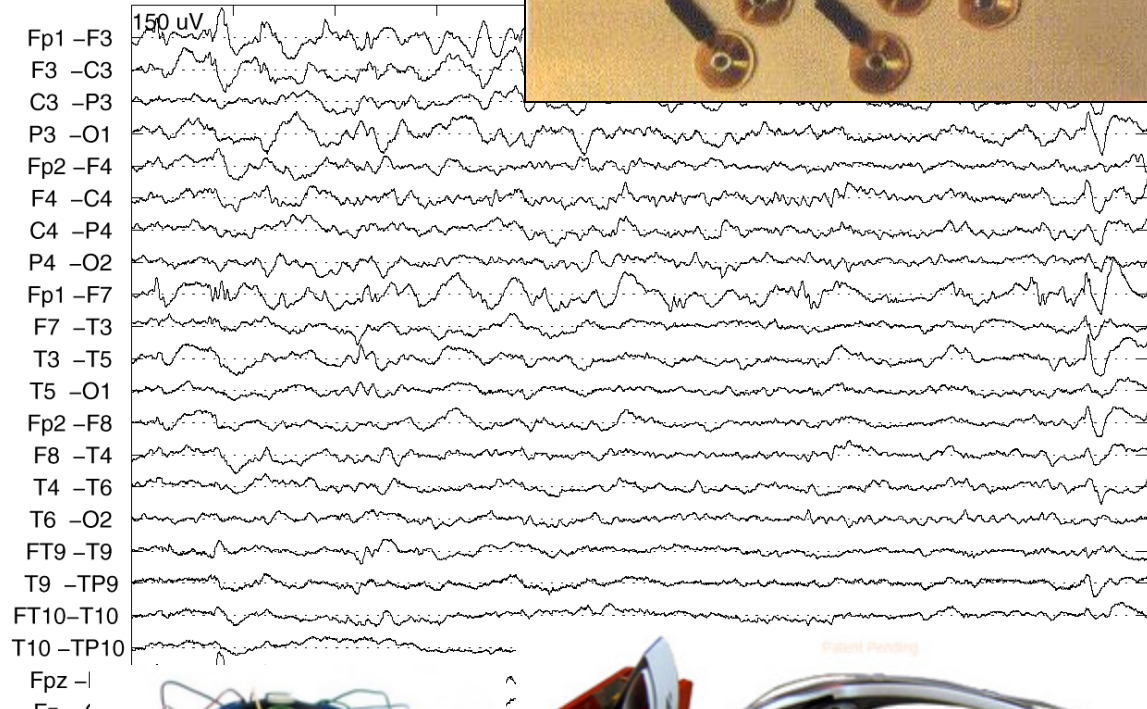
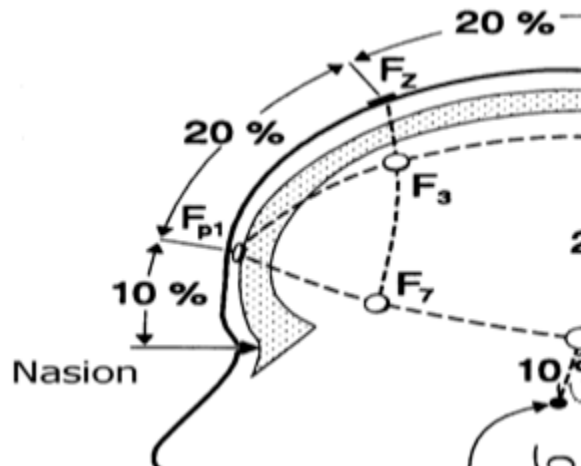
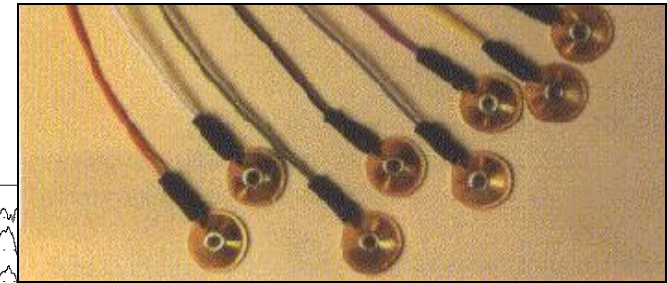




# Magnetoencephalography

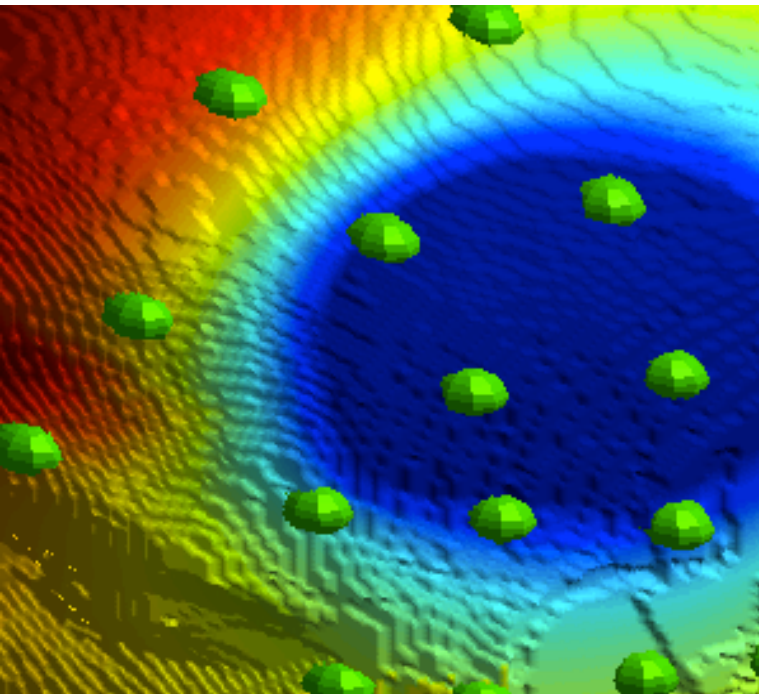


# Electroencephalography

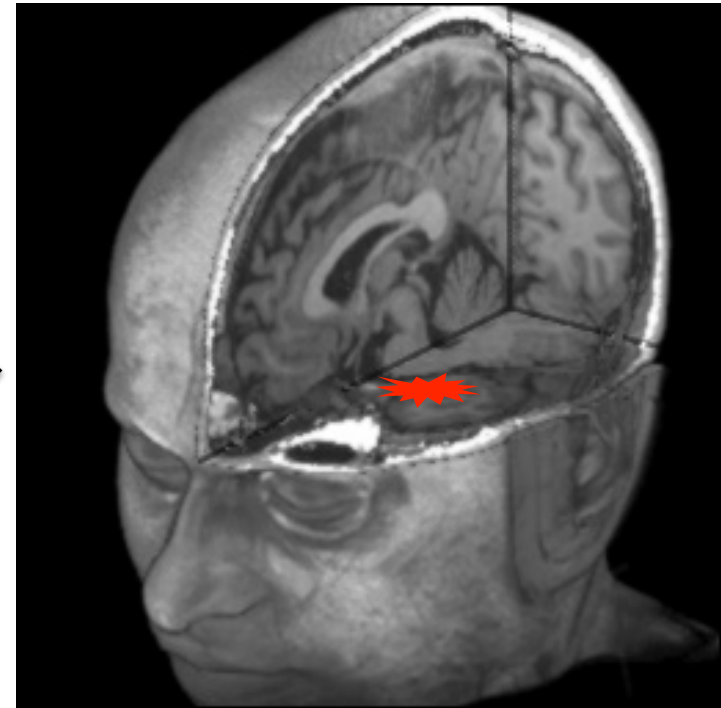


# EEG/MEG source localization

Potential distribution at the scalp  
measured at electrodes or sensors



Estimation of the origin of the  
electrical activity in the brain

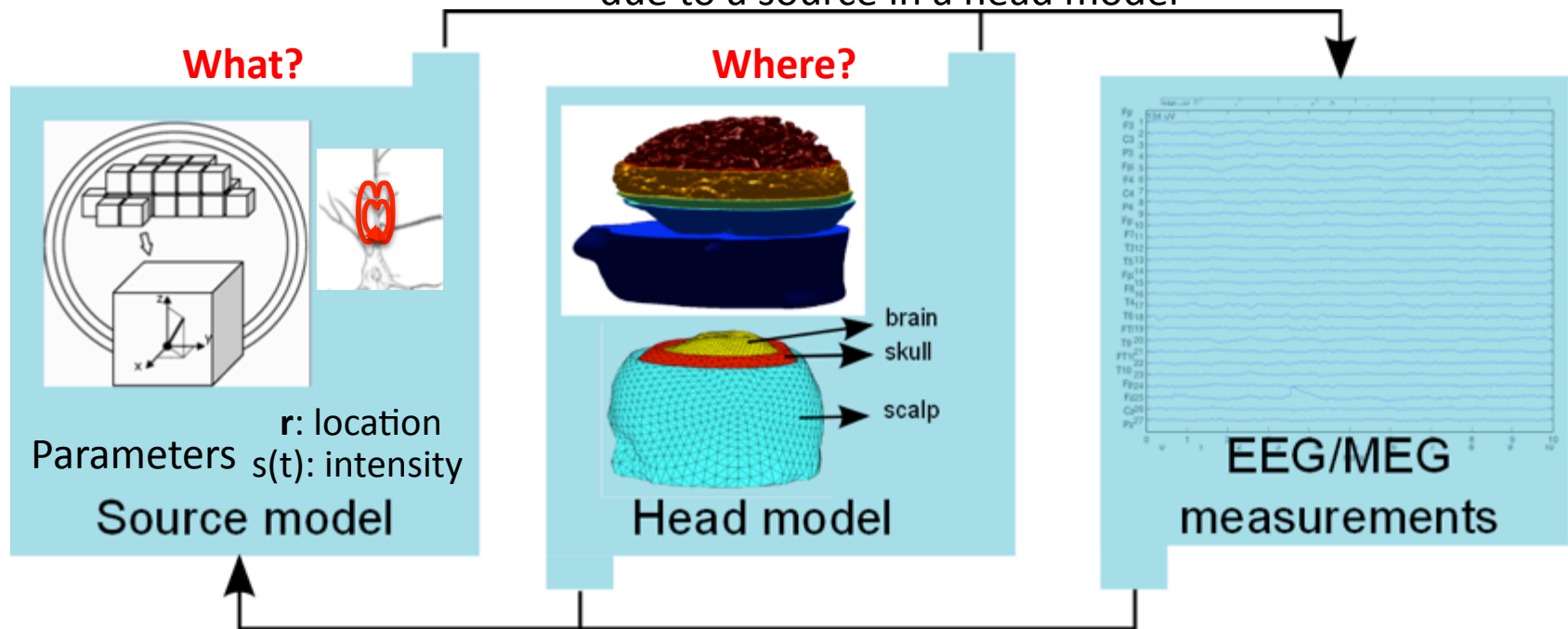




# EEG/MEG source localization

## Forward problem **How?**

Calculation of potentials or magnetic field  
due to a source in a head model



## Inverse problem **How?**

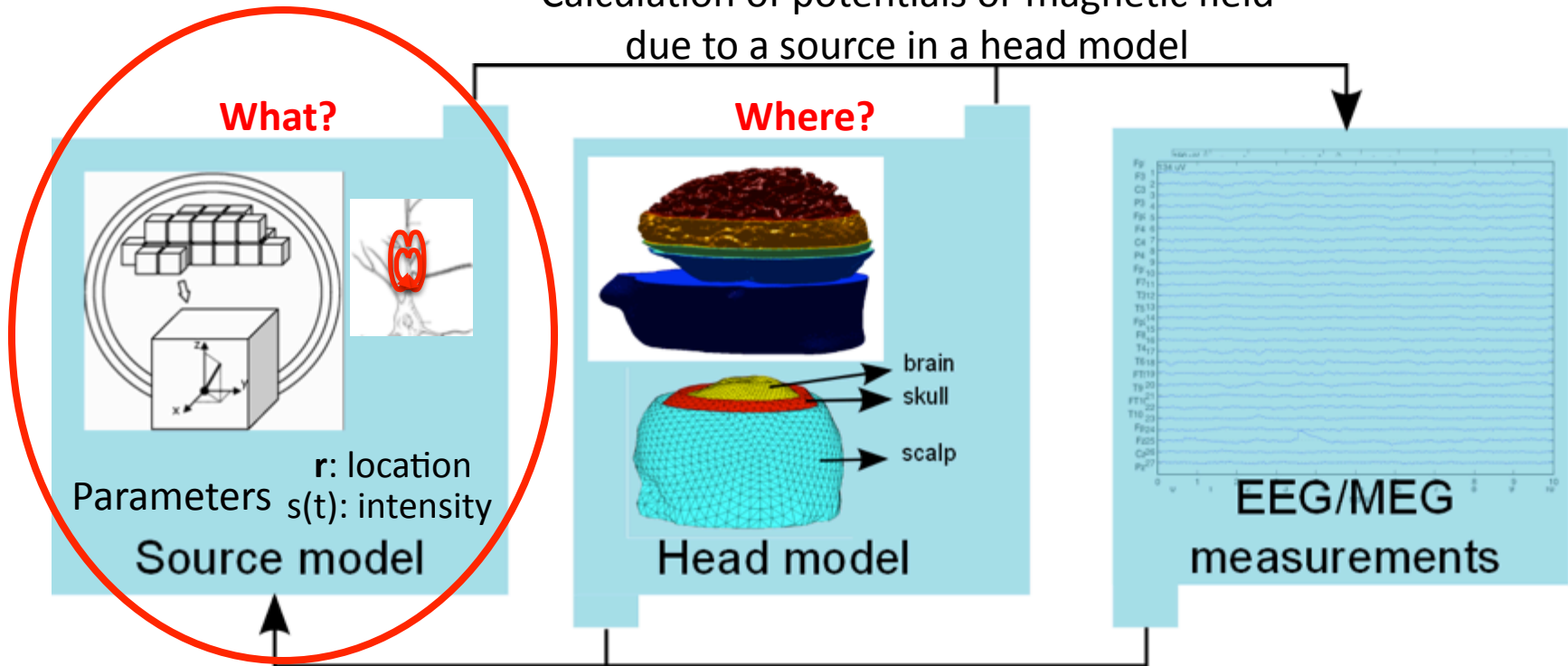
Estimation of the source parameters  
given the EEG/MEG and a head model



# EEG/MEG source localization

## Forward problem **How?**

Calculation of potentials or magnetic field  
due to a source in a head model

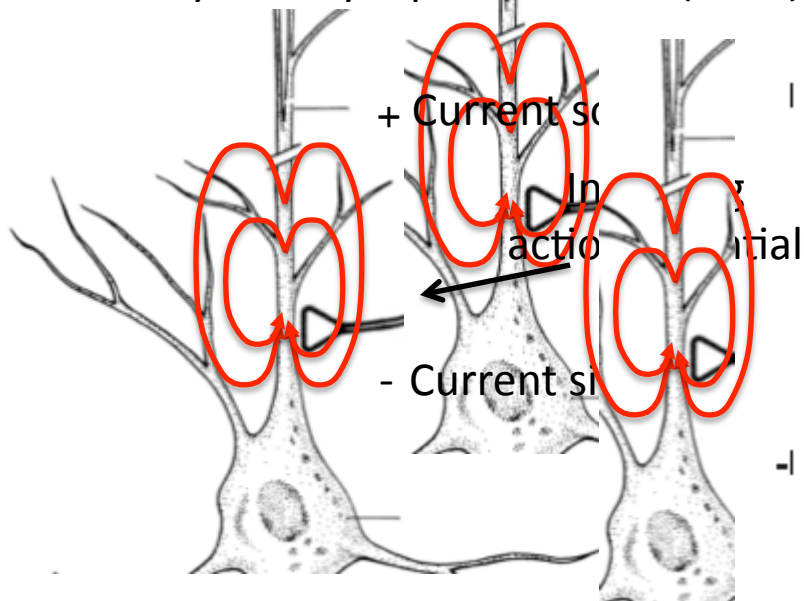


## Inverse problem **How?**

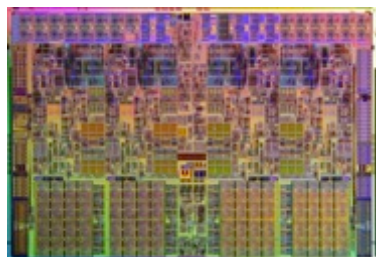
Estimation of the source parameters  
given the EEG/MEG and a head model

# Source model : generator EEG revisited

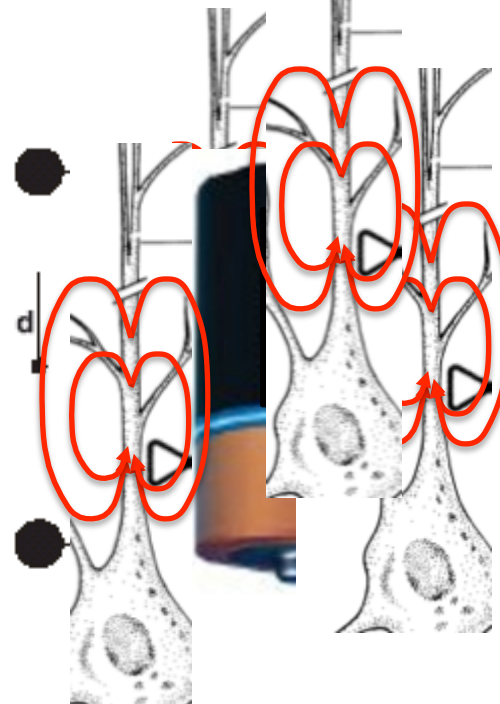
Excitatory Post-Synaptic Potential (EPSP)



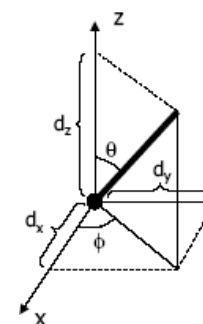
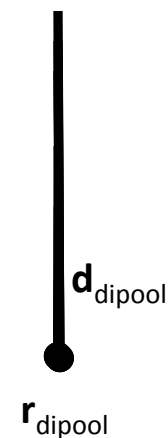
Microscopic current  
due to 1 EPSP :  $5 \cdot 10^{-5} \text{ nAm}$



100 times smaller than the current  
in a transistor on the Intel Core i7



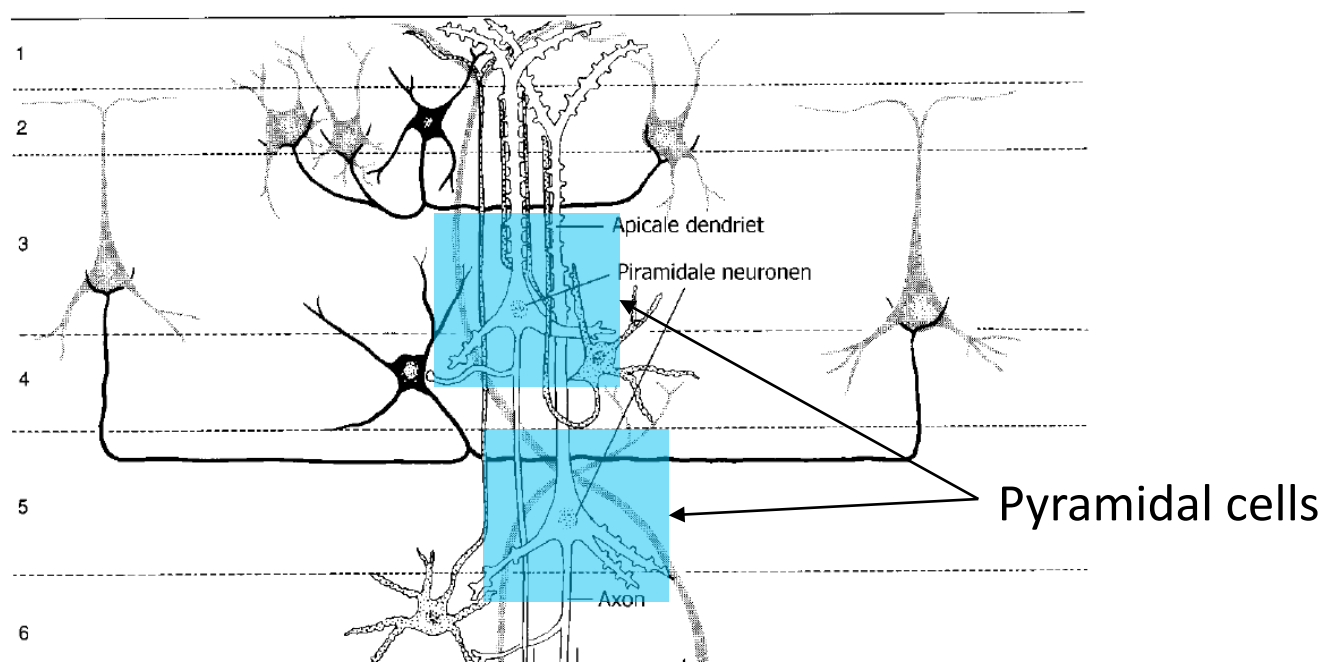
Macroscopic current measured: **50 nAm**  
equivalent to a cortical patch of **5.5x5.5 mm<sup>2</sup>**



6 parameters  
• Location  $\mathbf{r} (x,y,z)$   
• Orientation :  $\theta, \phi$   
• Intensity :  $s$

# Source model : generators of EEG revisited

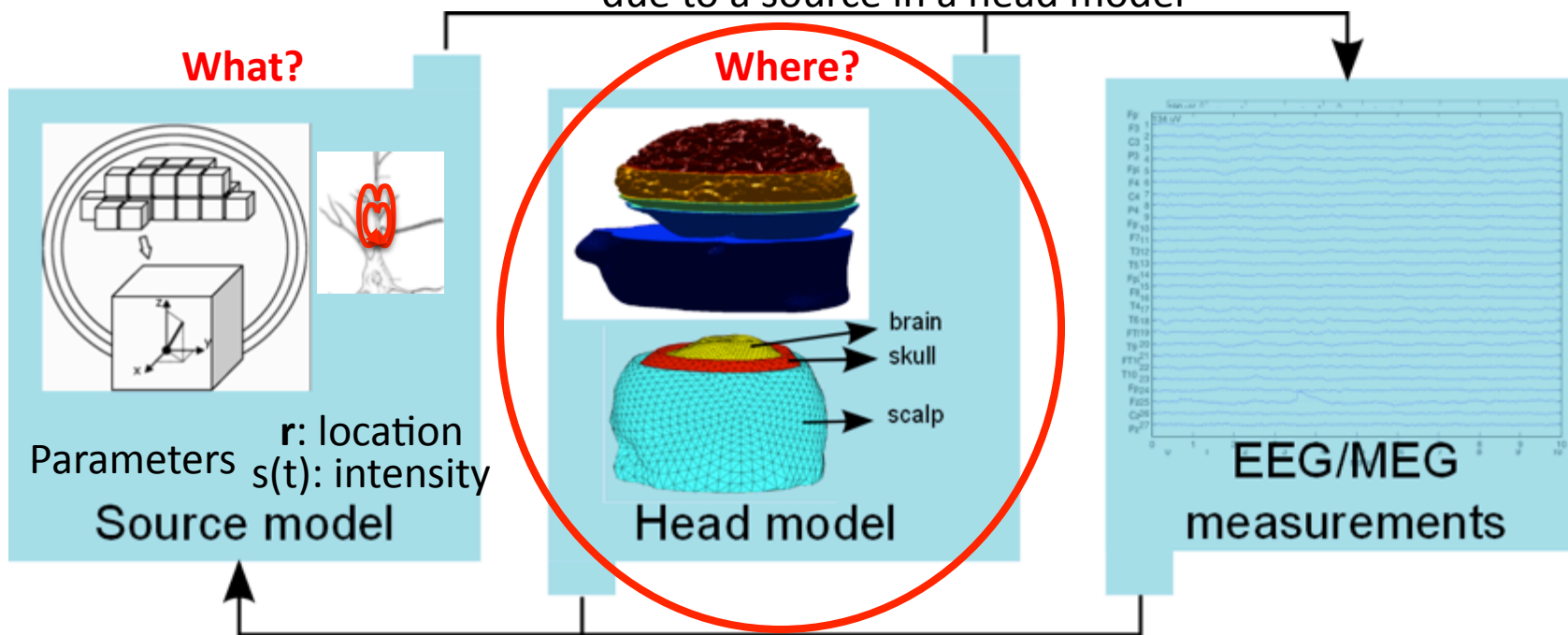
- Superposition of *large group of synchronously activated* neurons required to produce measurable potentials at the scalp
- Pyramidal neurons
  - Long apical dendrites oriented in parallel, perpendicular to the cortex
  - Are believed to be the main EEG generators



# EEG/MEG source localization

## Forward problem **How?**

Calculation of potentials or magnetic field  
due to a source in a head model



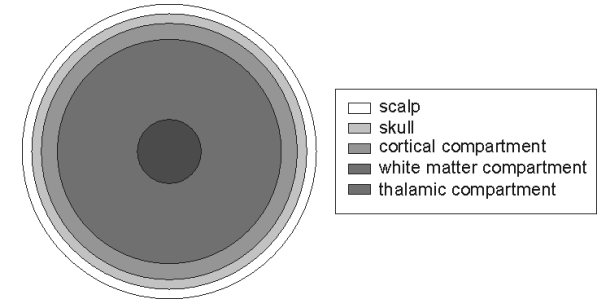
## Inverse problem **How?**

Estimation of the source parameters  
given the EEG/MEG and a head model

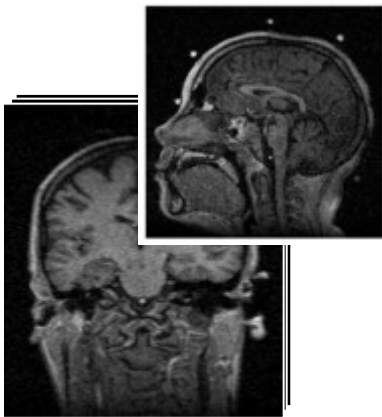


# Volume conductor modeling

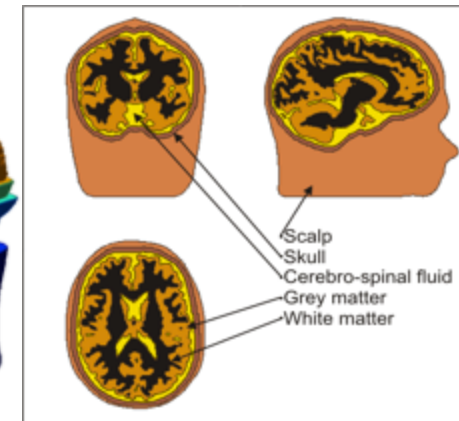
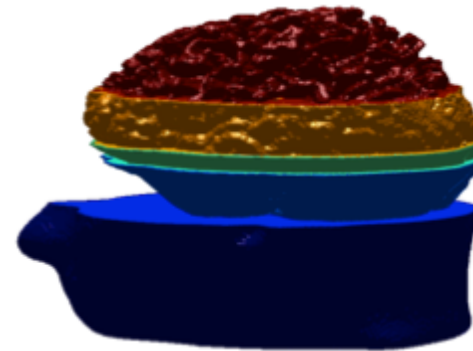
- Simplified spherical head models
  - Easy solution for forward problem



- Realistic head models



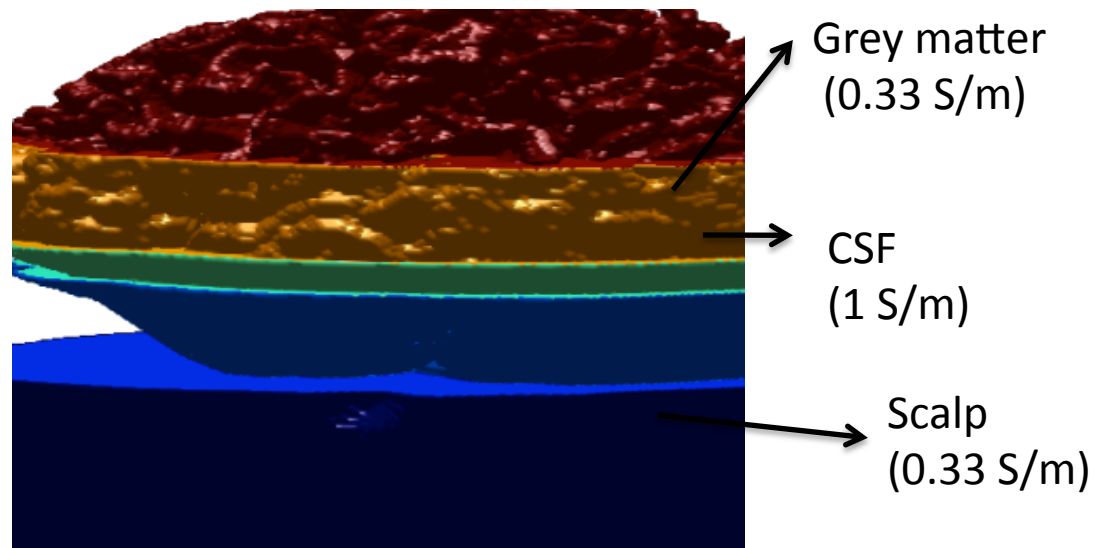
Segmentation



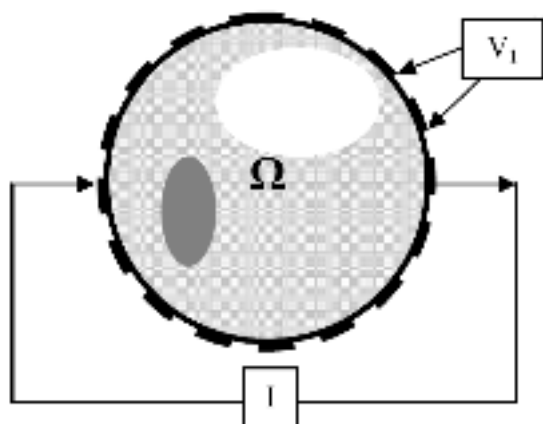
- Conductivities?

# Volume conductor modeling : conductivities

- Large uncertainty
  - Isotropic conductivities



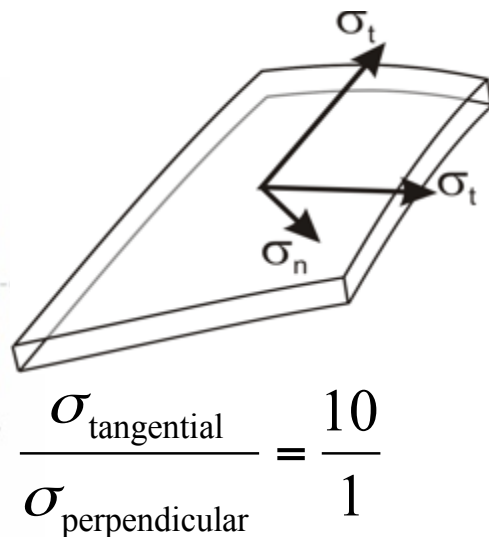
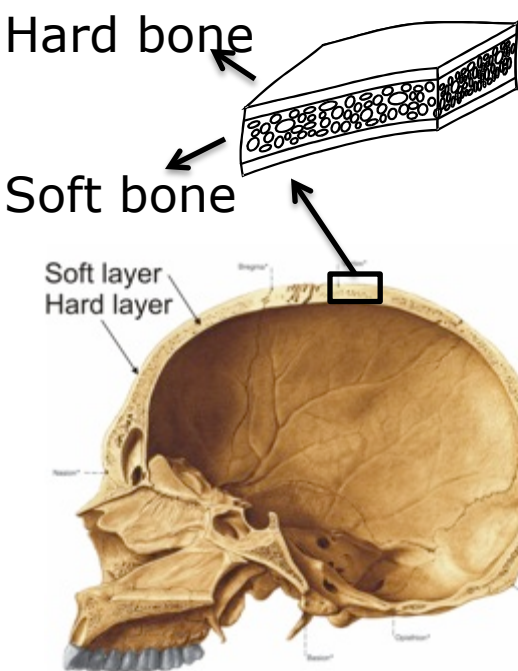
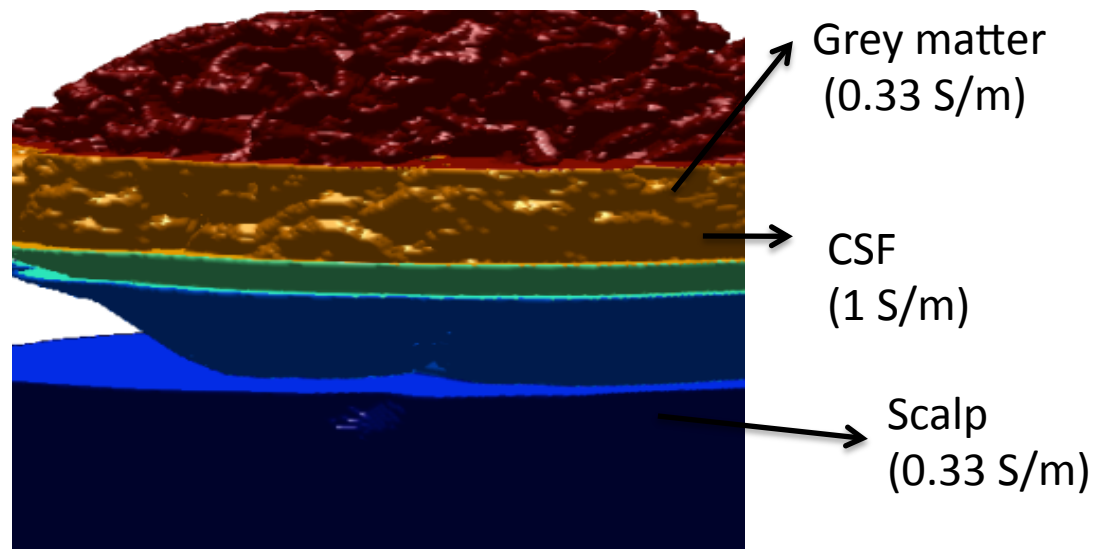
Electrical Impedance Tomography (EIT)



Magnetic Resonance - EIT

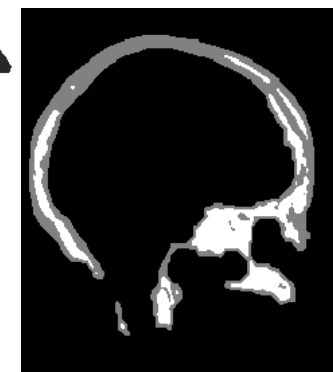
# Volume conductor modeling : conductivities

- Large uncertainty
  - Isotropic conductivities
  - Skull as anisotropic



Skull as a layered conductor

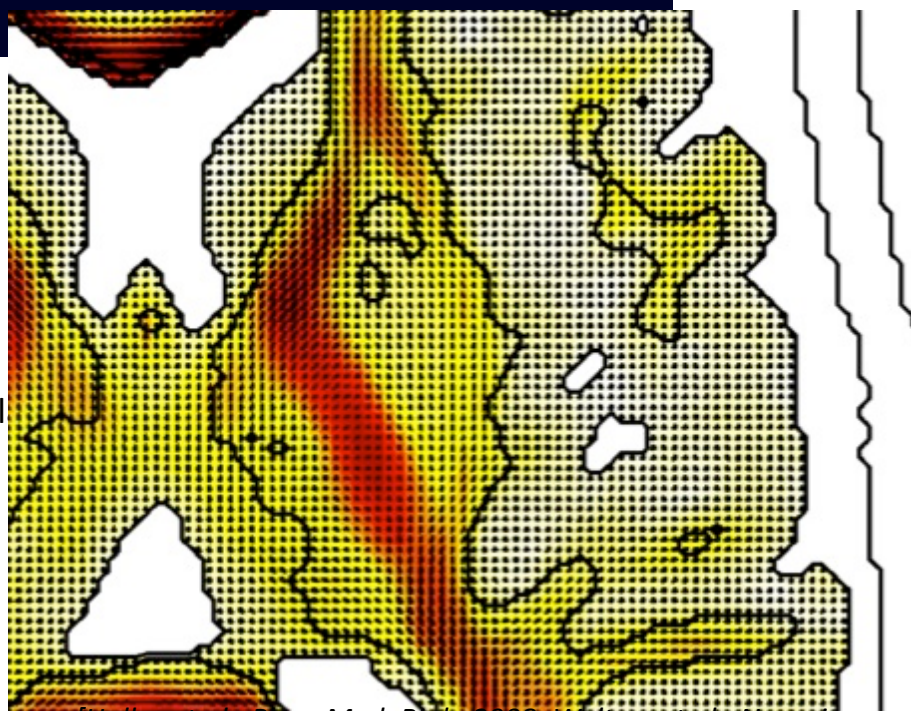
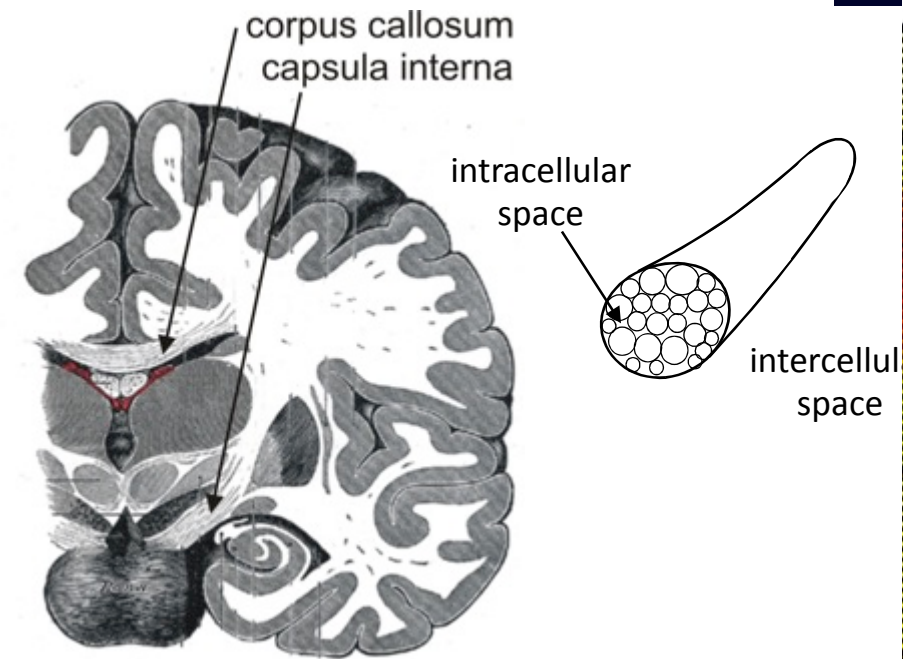
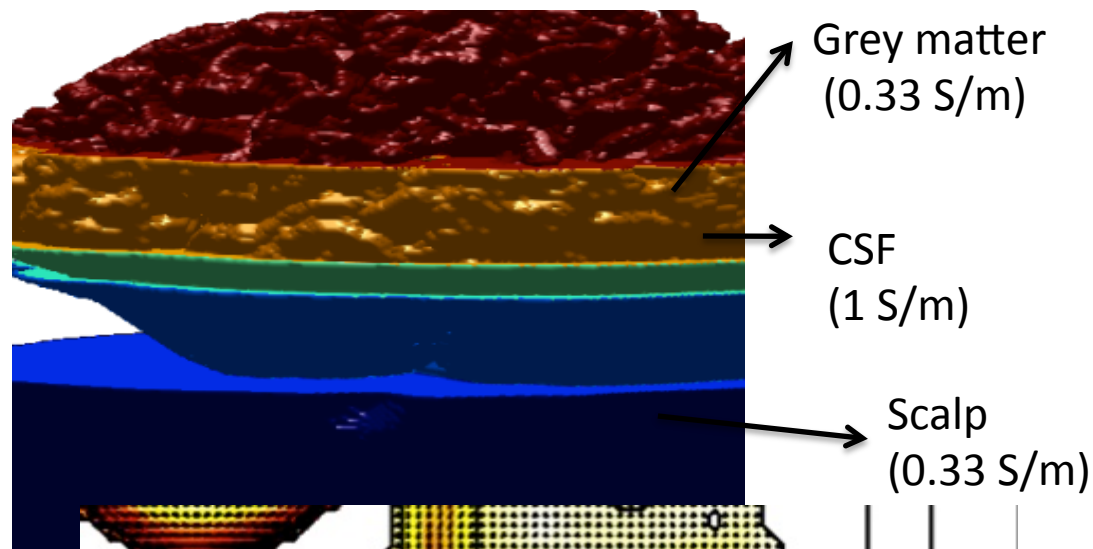
$$\frac{\sigma_{\text{tangential}}}{\sigma_{\text{perpendicular}}} = 1.8$$



[Montes et al., in preparation]

# Volume conductor modeling : conductivities

- Large uncertainty
  - Isotropic conductivities
  - Skull as anisotropic
  - White matter nerve bundles as anisotropic



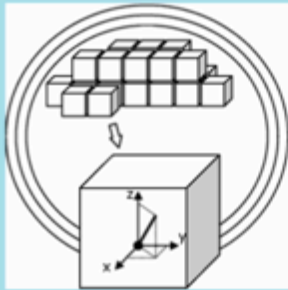


# EEG/MEG source localization

## Forward problem **How?**

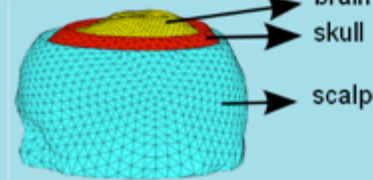
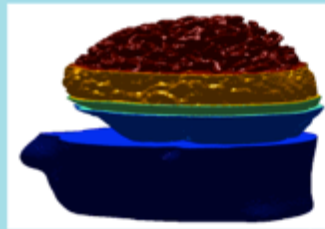
Calculation of potentials or magnetic field due to a source in a head model

**What?**

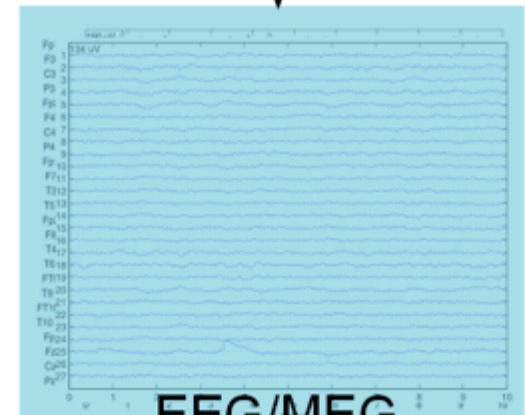


Parameters  $r$ : location  
 $s(t)$ : intensity  
**Source model**

**Where?**



**Head model**



**EEG/MEG measurements**

## Inverse problem **How?**

Estimation of the source parameters given the EEG/MEG and a head model

# Forward problem

- Calculation of the electrode potentials given the head model, electrode positions and the source

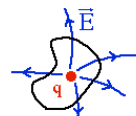
- Quasi-static approach

$$\nabla \cdot (\sigma(x, y, z) \cdot \nabla V(x, y, z)) = \nabla \cdot \mathbf{J}(x, y, z)$$

## Maxwell's Equations


$$\oiint \mathbf{E} \cdot \mathbf{n} \, dS = \frac{q}{\epsilon_0}$$

Gauss's Law



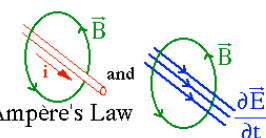
$$\oiint \mathbf{B} \cdot \mathbf{n} \, dS = 0$$

(no monopoles)



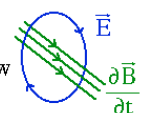
$$\oint \mathbf{B} \cdot d\mathbf{l} = \mu_0 \left( \mathbf{i} + \epsilon_0 \frac{d\Phi_E}{dt} \right)$$

Ampère's Law



$$\oint \mathbf{E} \cdot d\mathbf{l} = -\frac{d\Phi_B}{dt}$$

Faraday's Law



$$\nabla \cdot \mathbf{E} = \frac{\rho}{\epsilon_0} \quad \nabla \times \mathbf{B} = \mu_0 \left( \mathbf{j} + \epsilon_0 \frac{\partial \mathbf{E}}{\partial t} \right)$$

$$\nabla \cdot \mathbf{B} = 0 \quad \nabla \times \mathbf{E} = -\frac{\partial \mathbf{B}}{\partial t}$$

(Differential Forms)

Geometry

Electrode positions

Dipole parameters

Conductivities

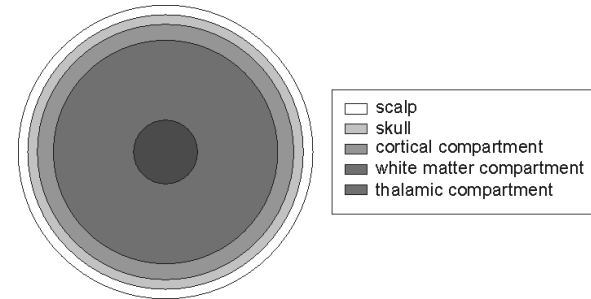


Potential at the electrodes



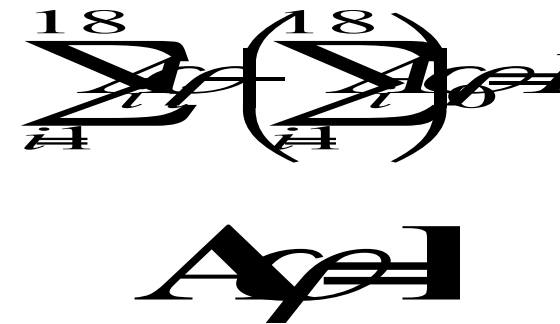
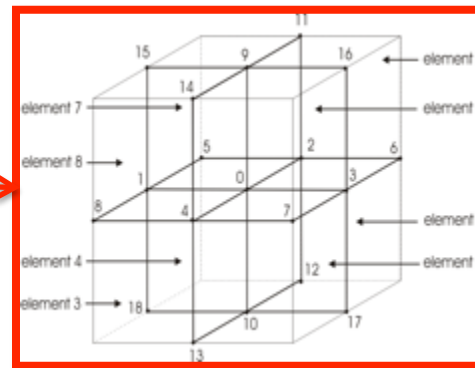
# Forward Problem

- Head model
  - Spherical head models
    - Analytical solution
    - Fast, but not realistic
  - Realistic head models
    - Numerical solutions
    - BEM, FEM, FDM

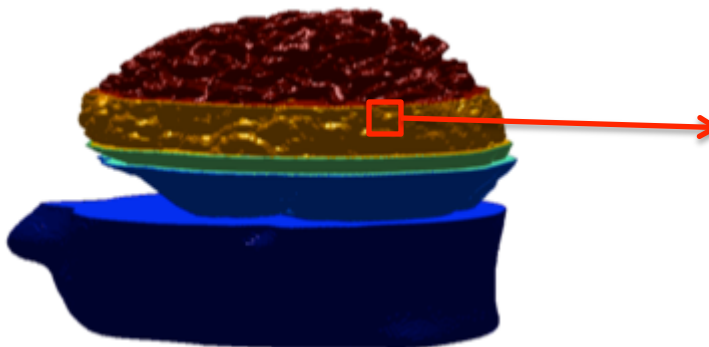


$$V(\mathbf{r}_{dip}, \mathbf{d}, \mathbf{r}_e) = \frac{d}{4\pi\xi_N r_e^2} \sum_{n=1}^{\infty} \frac{2n+1}{n} \left( \frac{r_{dip}}{r_e} \right)^{n-1} (f_n n \cos \alpha P_n(\cos \gamma) + g_n \cos \beta \sin \alpha P_n^1(\cos \gamma))$$

Potentials  $V$  are discretized into  $\varphi$

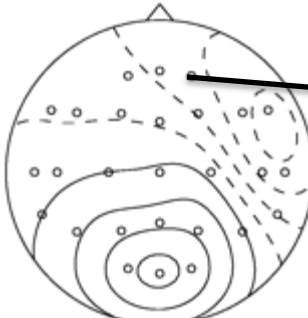


Linear system of equations of > 5 million equations



# Forward Problem : spatio-temporal model

- Linearity of the Maxwell equations



Intensity

$$v_{el}(t) = L(\mathbf{r}_d, \mathbf{r}_{el}) \mathbf{e}_d s(t) = a(\mathbf{r}_d, \mathbf{r}_{el}, \mathbf{e}_d) s(t)$$

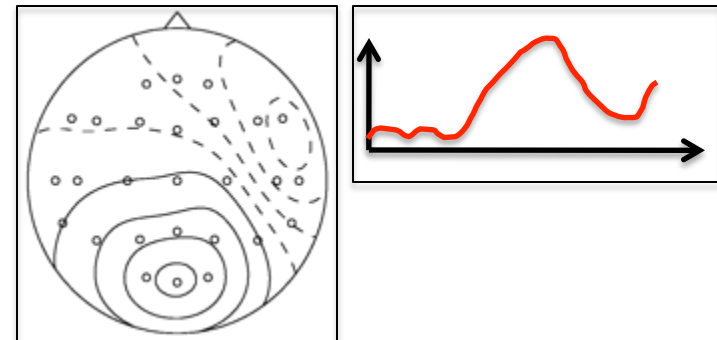
Lead-field matrix  
non-linear wrt  
dipole location

orientation

Contribution of a  
unitary dipole

Topography = potential distribution at the scalp on  $p$  electrodes

$$\mathbf{v}(t) = \begin{bmatrix} v_{el,1}(t) \\ \vdots \\ v_{el,p}(t) \end{bmatrix} = \begin{bmatrix} a(\mathbf{r}_d, \mathbf{r}_{el,1}, \mathbf{e}_d) \\ \vdots \\ a(\mathbf{r}_d, \mathbf{r}_{el,p}, \mathbf{e}_d) \end{bmatrix} s(t) = \mathbf{a}(\mathbf{r}_d, \mathbf{e}_d) \cdot s(t)$$

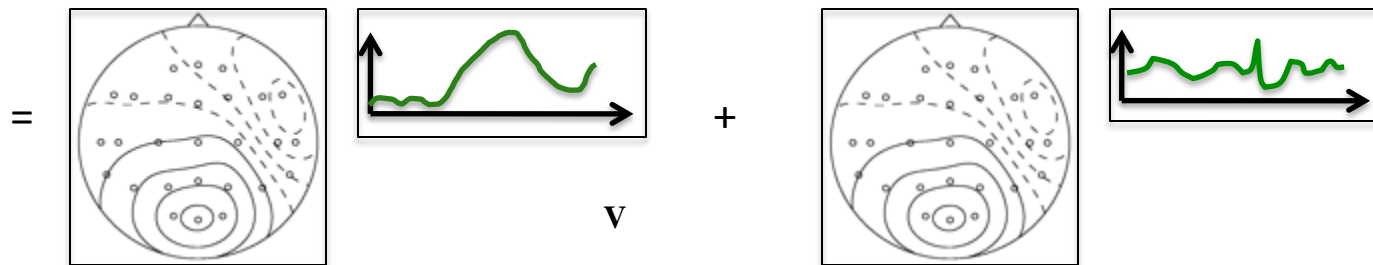
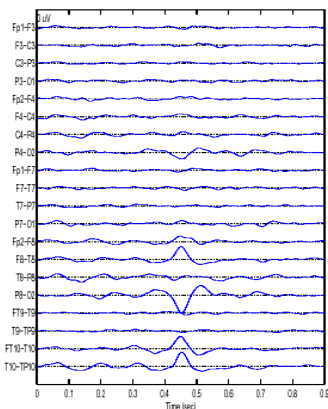




# Forward Problem : spatio-temporal model

- Linearity of the Maxwell equations

$$\mathbf{v}(t) = \mathbf{a}_1(\mathbf{r}_d, \mathbf{e}_d) \cdot s_1(t) + \mathbf{a}_2(\mathbf{r}_d, \mathbf{e}_d) \cdot s_2(t) + \dots$$



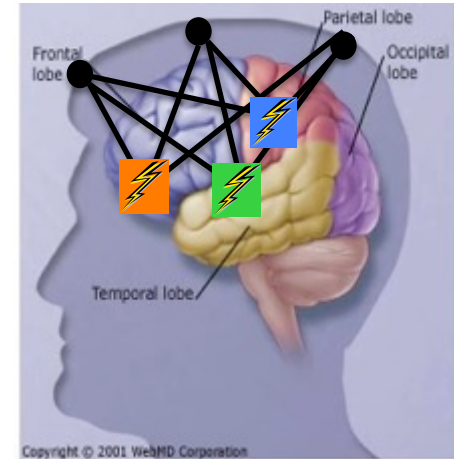
Spatio – temporal model

$$\mathbf{V} = \underbrace{\begin{bmatrix} \mathbf{a}_1(\mathbf{r}_d, \mathbf{e}_d) & \dots & \mathbf{a}_p(\mathbf{r}_d, \mathbf{e}_d) \end{bmatrix}}_{\text{Topographies}} \underbrace{\begin{bmatrix} s_1(t) \\ \vdots \\ s_p(t) \end{bmatrix}}_{\text{Source signals}} = \mathbf{A}\mathbf{S}^T$$

# Intermezzo : decomposition methods

Spatio – temporal model

$$\mathbf{V} = \underbrace{\begin{bmatrix} \mathbf{a}_1(\mathbf{r}_d, \mathbf{e}_d) & \dots & \mathbf{a}_p(\mathbf{r}_d, \mathbf{e}_d) \end{bmatrix}}_{\text{Topographies}} \underbrace{\begin{bmatrix} \mathbf{s}_1(t) \\ \vdots \\ \mathbf{s}_p(t) \end{bmatrix}}_{\text{Source signals}} = \mathbf{A}\mathbf{S}^T$$



Decomposition by  $p$  dipole sources

How to determine these:  
See inverse problem



- Neurophysiological basis
- Model-driven: controllable



- Model driven : many a priori known parameters
- Number of sources

Source signals

- Decomposition by  $m$  signal sources
- Based on properties of time series
  - Blind source separation
  - PCA, ICA, CCA



- Data-driven : no need for a complicated head model
- Few parameters

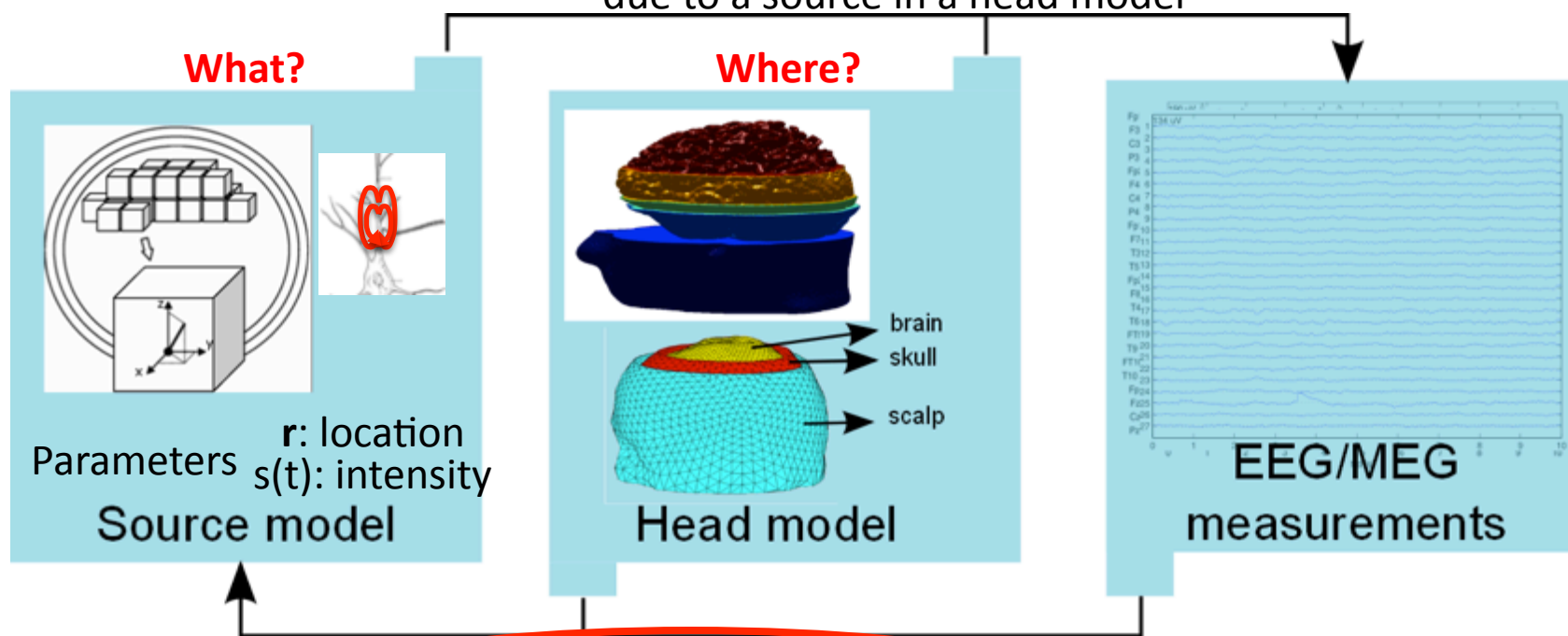


- Largely dependent on the data
- Sources should be stationary

# EEG/MEG source localization

## Forward problem **How?**

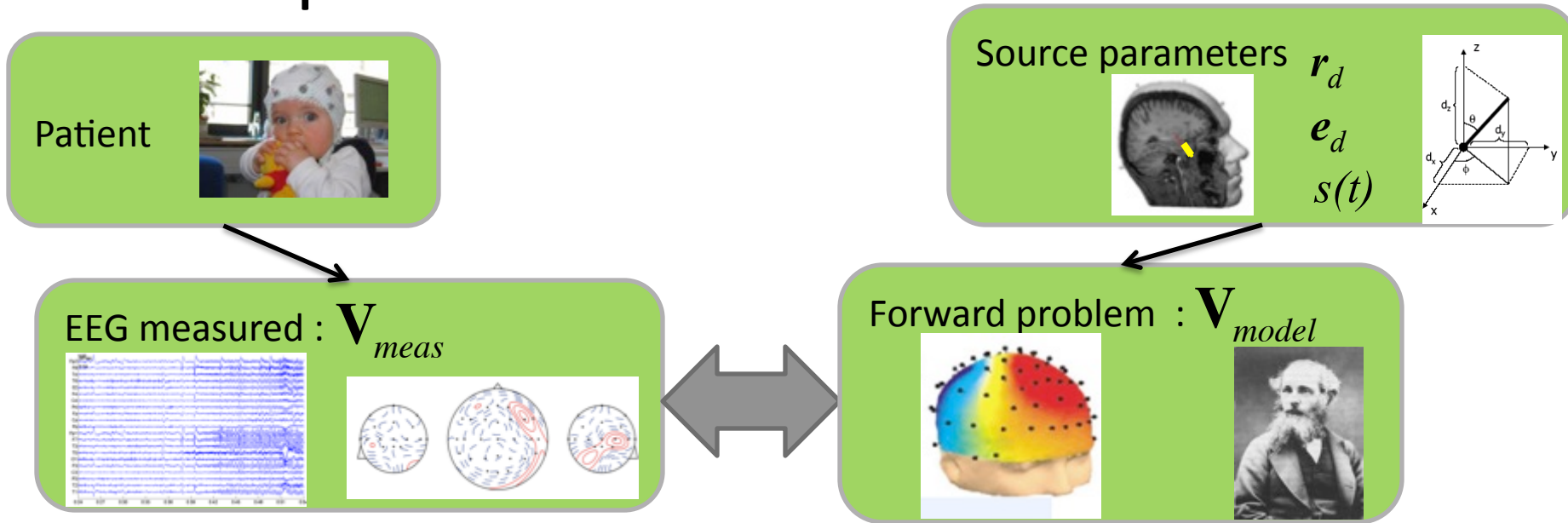
Calculation of potentials or magnetic field  
due to a source in a head model



## Inverse problem **How?**

Estimation of the source parameters  
given the EEG/MEG and a head model!

# Inverse problem



Estimation of the parameters by means of minimization of a cost function

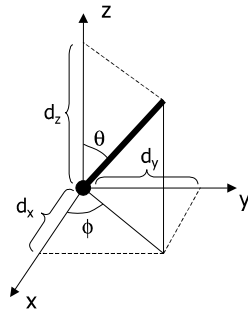
$$\begin{aligned}
 RE &= \left\| \mathbf{V}_{meas} - \mathbf{V}_{model} \right\| \\
 &\quad \downarrow \text{Spatio-temporal model} \\
 &= \left\| \mathbf{V}_{meas} - \mathbf{A}\mathbf{S}^T \right\| \quad ? \text{ Number of sources ?}
 \end{aligned}$$



# Inverse problem

$$RE = \left\| \mathbf{V}_{meas} - \mathbf{A}\mathbf{S}^T \right\|$$

**Assumption** : Within the EEG time frame there is only one source



**Single dipole model**

- Location
- Orientation
- Intensity

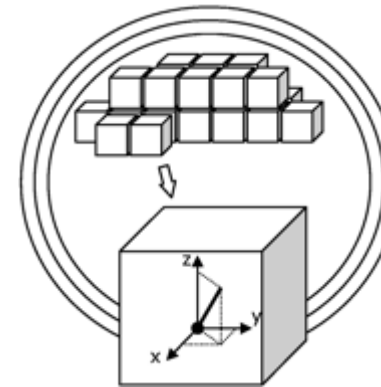
**Overdetermined system**

- #parameters (=6) < #electrodes

**Least squares solution**

- Best fit of the model onto the measurements
- Time consuming

**Assumption** : there are many sources



**Distributed source model**

- $\approx 10000$  dipoles distributed of grey matter
- Intensity

**Underdetermined system (ill-posed)**

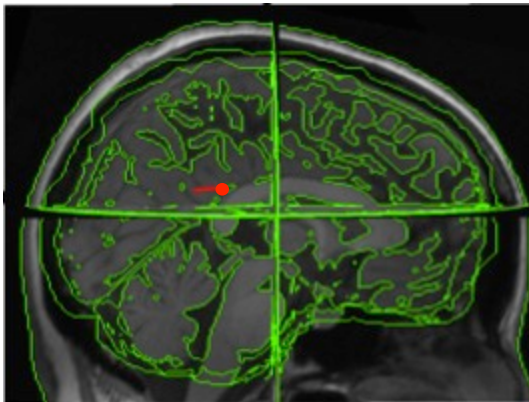
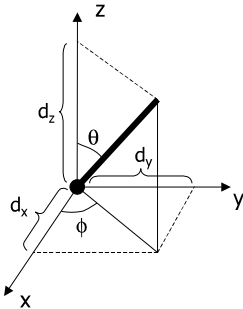
- #parameters > #electrodes

**Minimum norm solution**

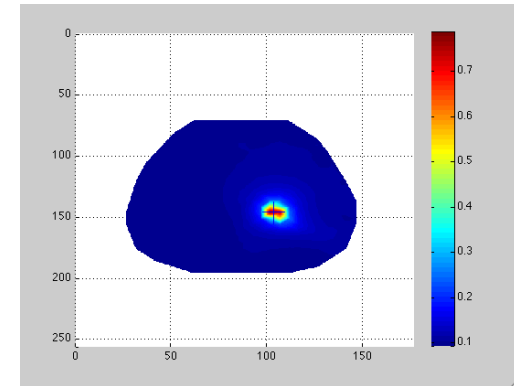
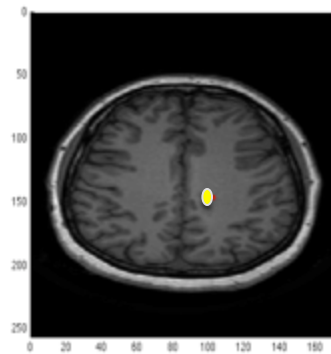
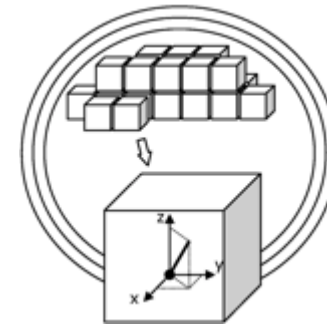
- Fast
- Regularization

# Inverse problem

$$RE = \left\| \mathbf{V}_{meas} - \mathbf{A}\mathbf{S}^T \right\|$$



Parametric solution



Imaging solution  
MNE, (s)LORETA, FOCUSS,  
beamformers

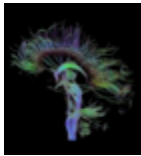
MUSIC, RAP – MUSIC, POP – MUSIC  
Multiple dipole localization (2 – 6 dipoles)

# Application: Localization of the epileptogenic zone

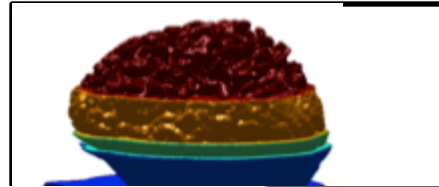
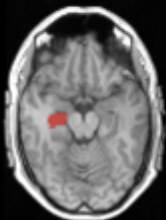
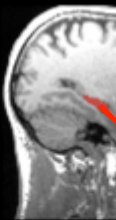
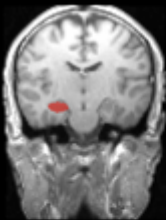
## Model



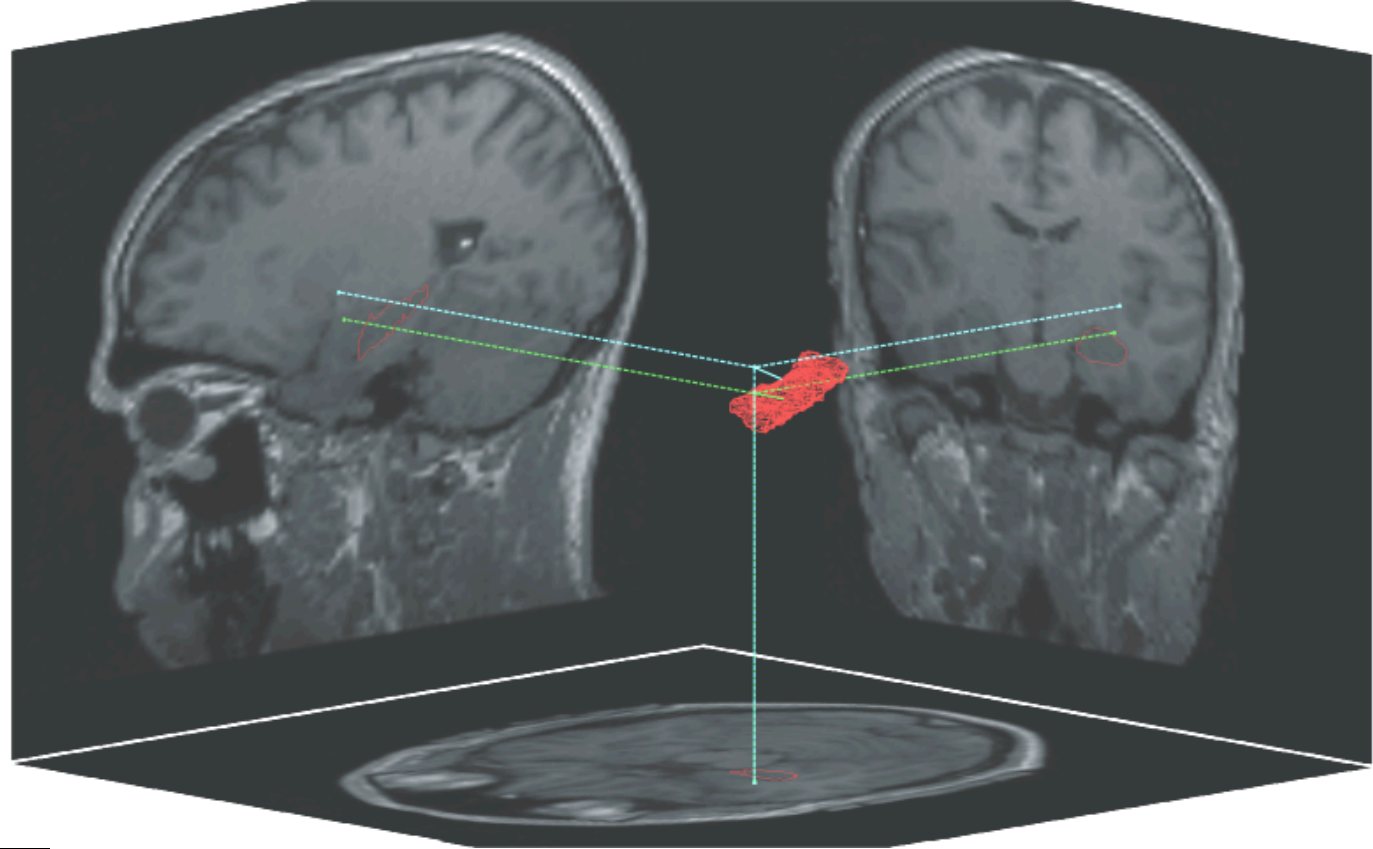
Traditional MRI scan



Diffusion weighted MRI



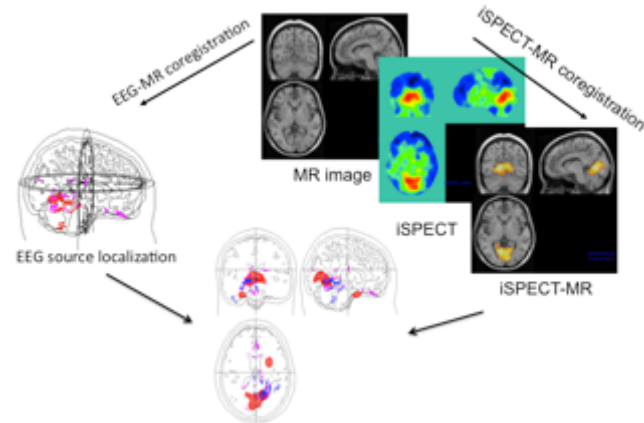
## Measurements Epileptic spike



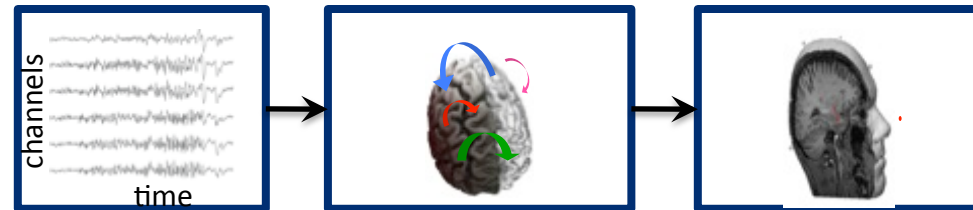
Estimation in head model with anisotropic conductivities

Estimation in head model with isotropic conductivities

## Multimodality



## Connectivity

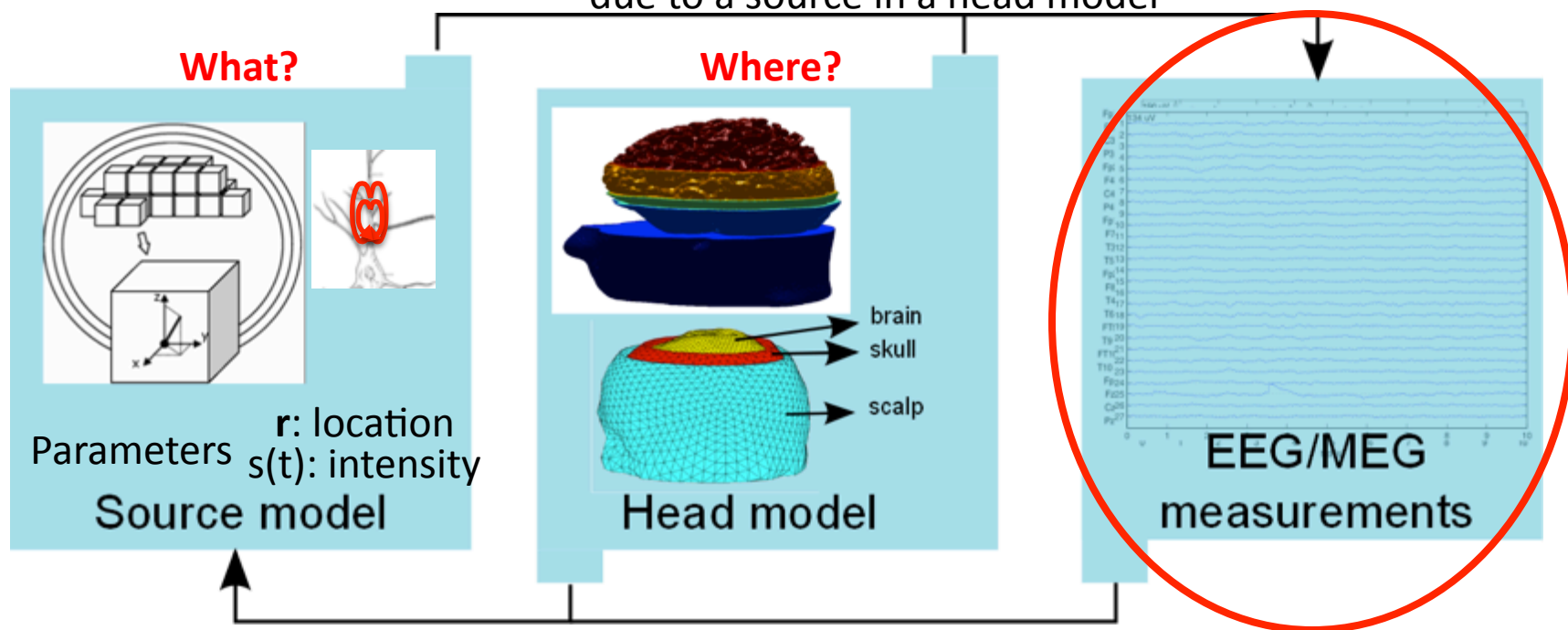


# RECENT TRENDS AND ADVANCES

# EEG/MEG source localization

## Forward problem **How?**

Calculation of potentials or magnetic field  
due to a source in a head model



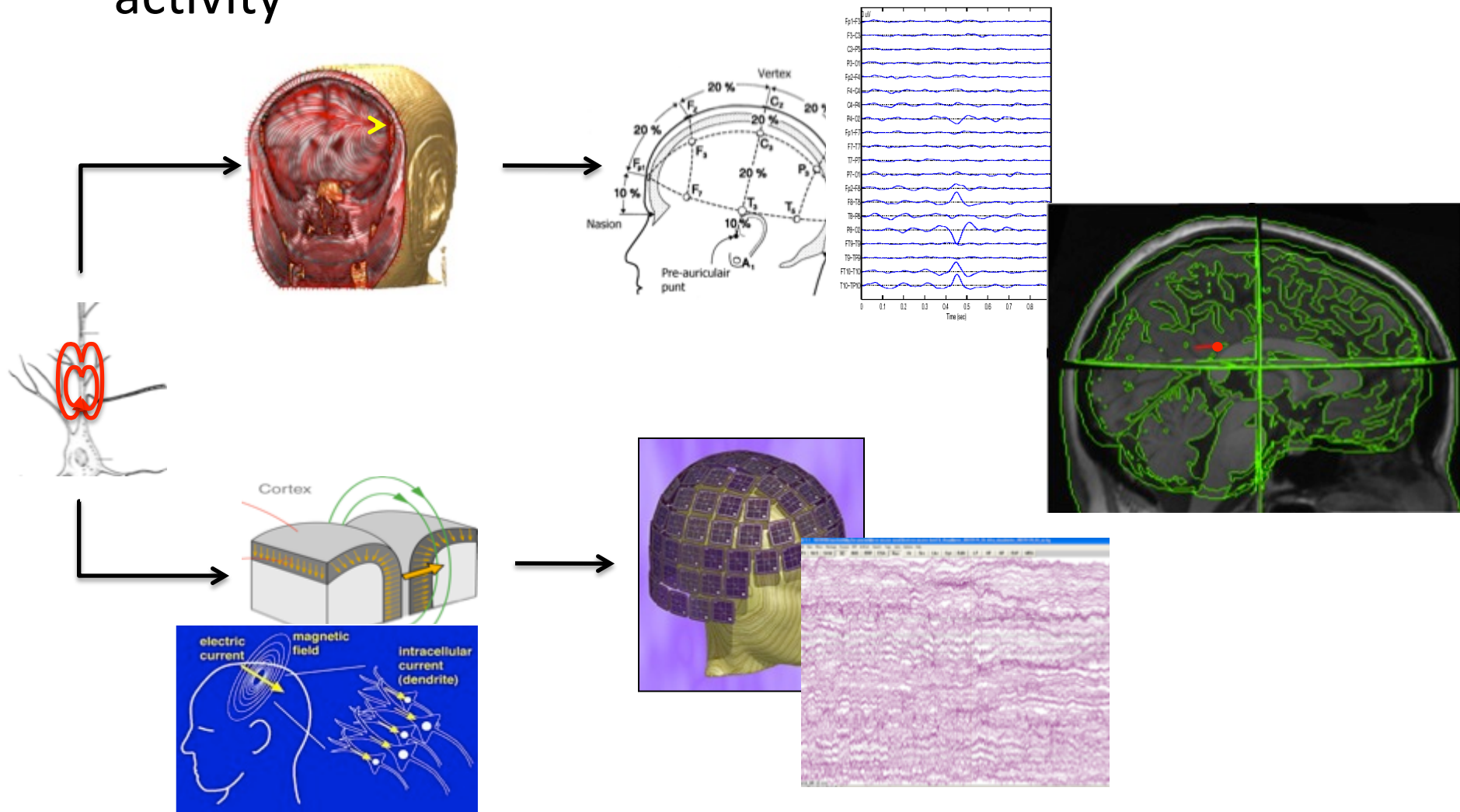
## Inverse problem **How?**

Estimation of the source parameters  
given the EEG/MEG and a head model



# Simultaneous EEG/MEG source localization

- EEG and MEG are generated by the same neuronal activity



# Multimodal integration of EEG source localization

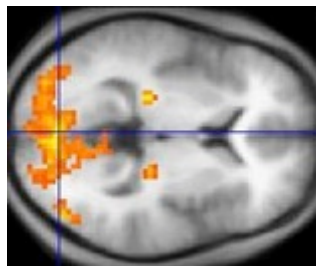
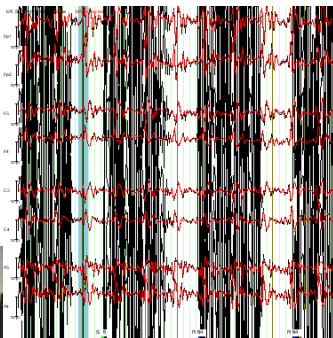
- Use functional imaging to improve spatial resolution of EEG source localization
  - Also provides timing information of the functional activation
  - A priori information of EEG source localization

Use functional MRI and EEG in epilepsy

Use ictal SPECT for localization of epileptic seizures

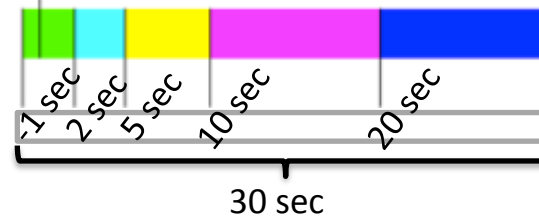
Preprocessing :

MR artefact  
BCG artefact

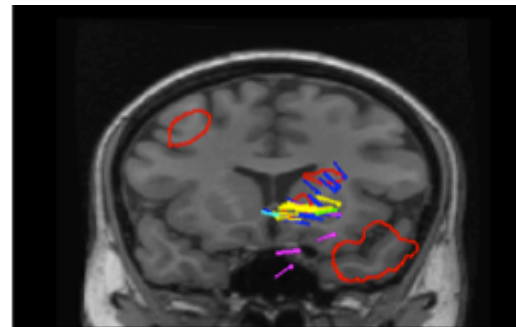


Time information of sources

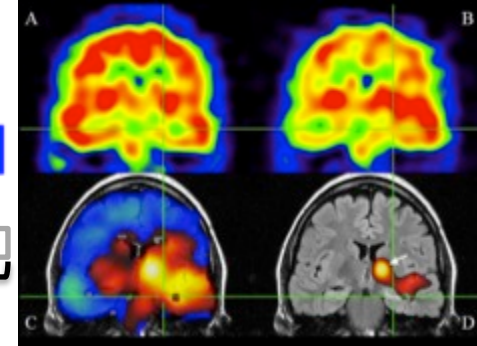
Start of seizure



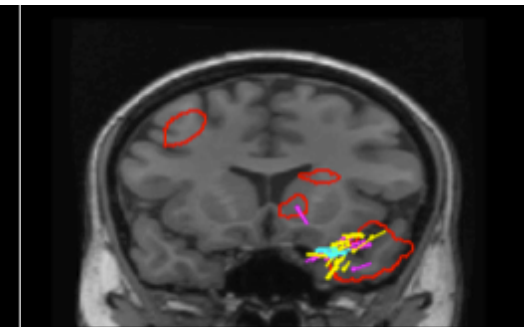
Without integration



Ictal SPECT



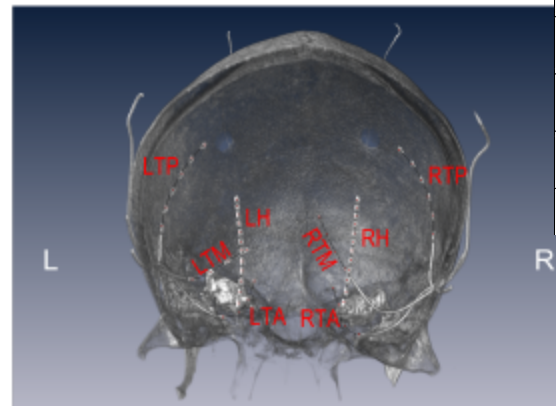
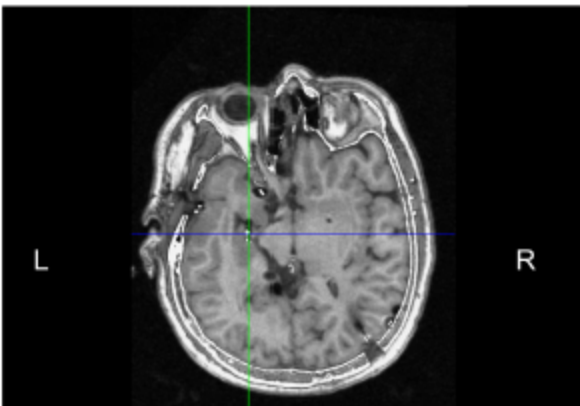
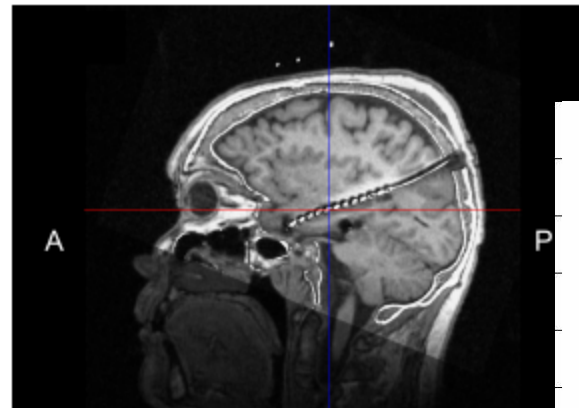
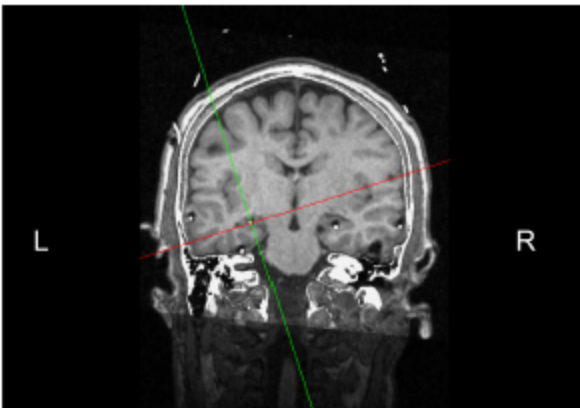
With integration



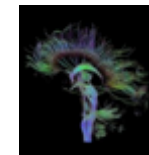
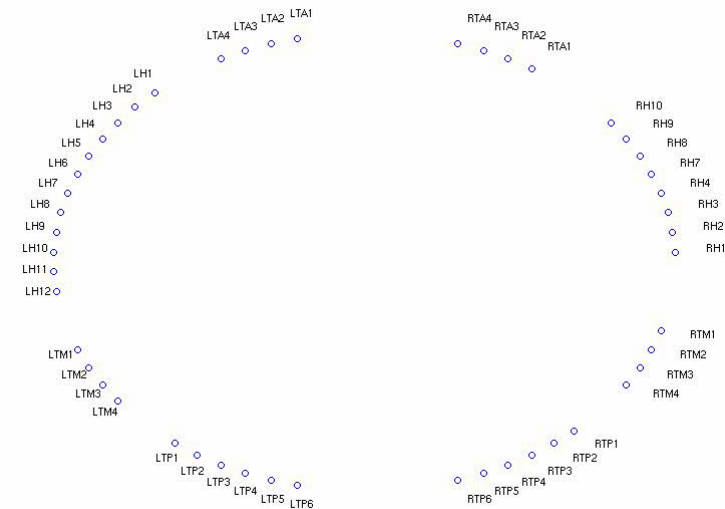
# Connectivity analysis

## Functional connectivity

propagation pattern of information flow between signals



Use of functional connectivity in to **accurately estimate to onset zone** of an epileptic seizure



Correlation with structural connectivity from fibretracking

[van Mierlo et al. 2010, Neuroimage, submitted]

# Conclusions

- EEG and MEG can be used to investigate the electrophysiological activations in the brain
  - Adequate model (source, head model)
  - Link with anatomy
- Trends
  - combination of multiple modalities
    - to increase the resolution
    - to exploit the time information
  - Connectivity
    - visualize flow of information

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# Thank you for your attention

